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10 30  
atgacgactgaaccggtattttttcaagcctggttttcaaagaaagaatt  
M T T E P L F F K P V F K E R I

50 70 90  
tggggcgggaccgcttttagctgattttggctataaccattccgtcacaa  
W G G T A L A D F G Y T I P S Q

110 130  
cgaacaggggagtgctgggcttttgccgcgcacataaaatgggtcaaagc  
R T G E C W A F A A H Q N G Q S

150 170 190  
gttggttcaaaacggaatgtataaggggttcacgctcagcgaattatgg  
V V Q N G M Y K G F T L S E L W

210 230  
gaacatcacagacattttatttcggacagcttgaaggggaccgtttccct  
E H H R H L F G Q L E G D R F P

250 270 2  
ctgcttacaaaaatattagatgctgaccaggacttatctgttcagggtg  
L L T K I L D A D Q D L S V Q V

90 310 330  
catccgaatgatgaatatgccaacatacatgaaaacgggtgagcttggg  
H P N D E Y A N I H E N G E L G

350 370  
aaaacagaatgctgggtacattattgattgccaaaaagatgccgagatt  
K T E C W Y I I D C Q K D A E I

390 410 430  
atttatggccacaatgcaacaacaaaggaagaactaactaccatgata  
I Y G H N A T T K E E L T T M I

450 470  
gagcgtggagaatgggatgagctcttgccgccgtgtaaagggtaaagccg  
E R G E W D E L L R R V K V K P

490 510 5  
ggggattttttctatgtgccaagcgggtactgttcattgcgattggaaaa  
G D F F Y V P S G T V H A I G K

30 550 570  
ggaattcttgctttggagacgcagcagaactcagacacaacctacaga  
G I L A L E T Q Q N S D T T Y R

FIG. 1A

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590 610  
ttatatgattatgaccgaaaagatgcagaaggcaagctgcgcgagctt  
L Y D Y D R K D A E G K L R E L

630 650 670  
catctgaaaaagagcattgaagtgatagagggtcccgtctattccagaa  
H L K K S I E V I E V P S I P E

690 710  
cggcatacagttcaccatgaacaaattgaggatttgcttacaacgaca  
R H T V H H E Q I E D L L T T T

730 750 7  
ttgattgaatgcgcttacttttcggtgggggaaatggaacttatcagga  
L I E C A Y F S V G K W N L S G

70 790 810  
tcagcaagcttaaagcagcaaaaaccattccttcttatcagtgatgatt  
S A S L K Q Q K P F L L I S V I

830 850  
gaaggggaggggccgtatgatctctggtgagtatgtctatcctttcaaa  
E G E G R M I S G E Y V Y P F K

870 890 910  
aaaggagatcatatgttgctgccttacgggtcttggagaatttaaactc  
K G D H M L L P Y G L G E F K L

930  
gaaggatatgcagaatgtatcgtctcccatctg  
E G Y A E C I V S H L

**FIG. 1B**

[illegible]

④  $\rightarrow$

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• •

[illegible]

Year	1990	1991	1992	1993	1994	1995	1996	1997	1998	1999	2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020	2021	2022	2023	2024	2025	2026	2027	2028	2029	2030	2031	2032	2033	2034	2035	2036	2037	2038	2039	2040	2041	2042	2043	2044	2045	2046	2047	2048	2049	2050	2051	2052	2053	2054	2055	2056	2057	2058	2059	2060	2061	2062	2063	2064	2065	2066	2067	2068	2069	2070	2071	2072	2073	2074	2075	2076	2077	2078	2079	2080	2081	2082	2083	2084	2085	2086	2087	2088	2089	2090	2091	2092	2093	2094	2095	2096	2097	2098	2099																																																																																																																																																																																																																										
1990	120	121	122	123	124	125	126	127	128	129	130	131	132	133	134	135	136	137	138	139	140	141	142	143	144	145	146	147	148	149	150	151	152	153	154	155	156	157	158	159	160	161	162	163	164	165	166	167	168	169	170	171	172	173	174	175	176	177	178	179	180	181	182	183	184	185	186	187	188	189	190	191	192	193	194	195	196	197	198	199	200	201	202	203	204	205	206	207	208	209	210	211	212	213	214	215	216	217	218	219	220	221	222	223	224	225	226	227	228	229	230	231	232	233	234	235	236	237	238	239	240	241	242	243	244	245	246	247	248	249	250	251	252	253	254	255	256	257	258	259	260	261	262	263	264	265	266	267	268	269	270	271	272	273	274	275	276	277	278	279	280	281	282	283	284	285	286	287	288	289	290	291	292	293	294	295	296	297	298	299	300	301	302	303	304	305	306	307	308	309	310	311	312	313	314	315	316	317	318	319	320	321	322	323	324	325	326	327	328	329	330	331	332	333	334	335	336	337	338	339	340	341	342	343	344	345	346	347	348	349	350	351	352	353	354	355	356	357	358	359	360	361	362	363	364	365	366	367	368	369	370	371	372	373	374	375	376	377	378	379	380	381	382	383	384	385	386	387	388	389	390	391	392	393	394	395	396	397	398	399	400	401	402	403	404	405	406	407	408	409	410	411	412	413	414	415	416	417	418	419	420	421	422	423	424	425	426	427	428	429	430	431	432	433	434	435	436	437	438	439	440	441	442	443	444	445	446	447

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VI

**FIG. 2**

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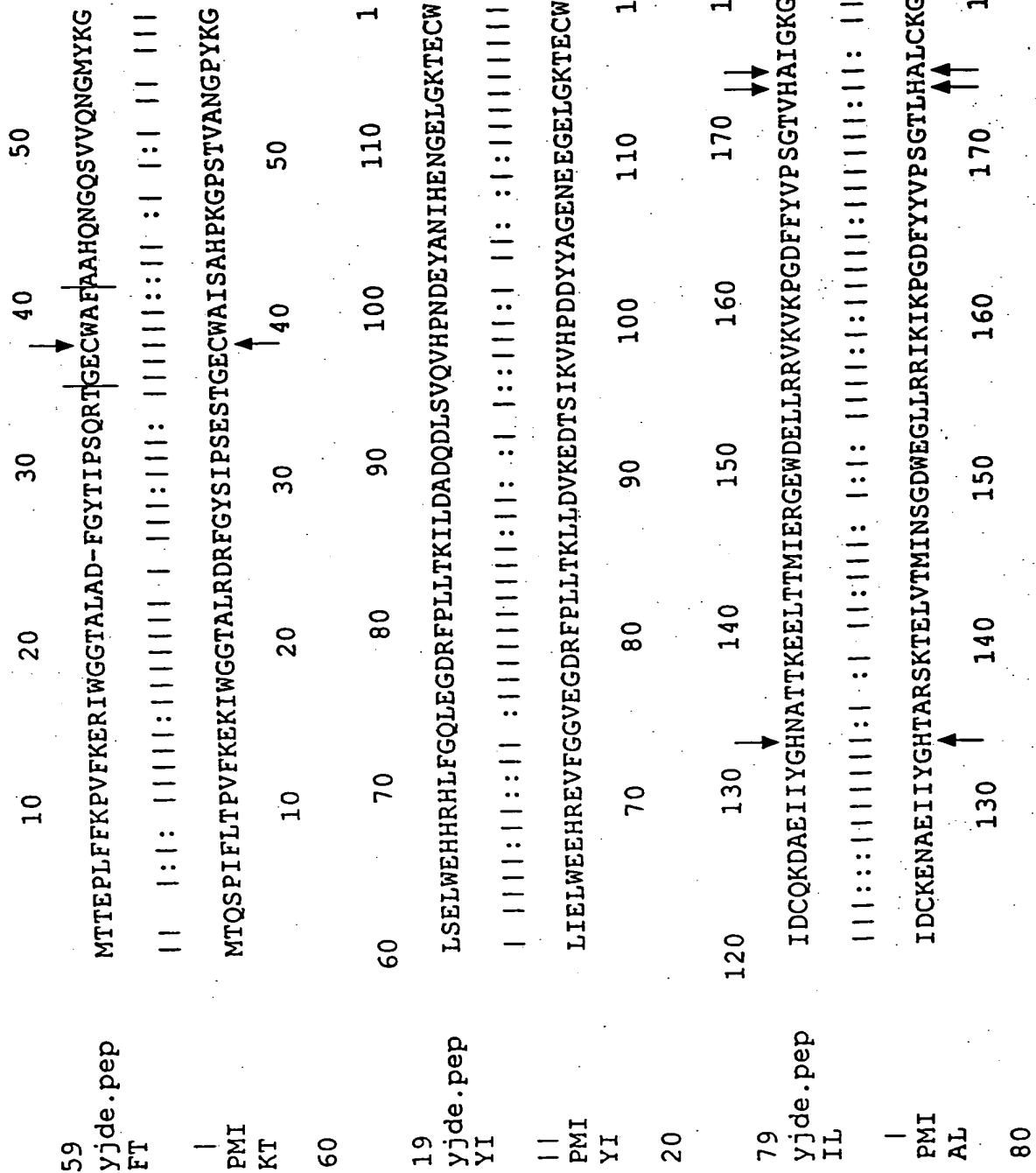


FIG.\_3A

**FIG. 3B**

```
yjde.pep      GEFKLEGYAEClVSHL
               :|::|  |||:
PMI           PDFTIKGTCTLIvSHI
               310
```

**FIG. 4A**

**FIG. 4B**

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10 30  
atgacgcattccattatTTTTtagagcctgtctTTTaaagaaagactatgg  
M T H P L F L E P V F K E R L W

50 70 90  
ggagggacgaagcttcgtgacgctTTTggctacgcaataccctcacaa  
G G T K L R D A F G Y A I P S Q

110 130  
aaaacaggtgagtgtggtggcgcttctgcacatgcccatggctcgctcg  
K T G E C W A V S A H A H G S S

150 170 190  
tctgtaaaaaatggcccgctggcaggaaagacacttgatcaagtatgg  
S V K N G P L A G K T L D Q V W

210 230  
aaagatcatccagagatattcgggtttccggatggtaagggtgtttccg  
K D H P E I F G F P D G K V F P

250 270 2  
ctgctggtaaaagctgctggacgccaatatggatctctccgtgcaagtc  
L L V K L L D A N M D L S V Q V

90 310 330  
catcctgatgatgattatgcaaaactgcacgaaaatggcgaccttggt  
H P D D D Y A K L H E N G D L G

350 370  
aaaacggagtgctggtatatcattgattgcaaagatgacgccgaacta  
K T E C W Y I I D C K D D A E L

390 410 430  
atTTTgggacatcatgcaagcacaaaggaagagttcaaacaacgaata  
I L G H H A S T K E E F K Q R I

450 470  
gaaagcgggtgattggaacgggctgctgaggcgaatcaaaatcaagcca  
E S G D W N G L L R R I K I K P

490 510 5  
ggagatttctTTTtatgtgccaaagcggtacactccatgctTTTatgtaag  
G D F F Y V P S G T L H A L C K

30 550 570  
ggaacccttgtccttgaaatccagcaaaactctgatacaacatatcgc  
G T L V L E I Q Q N S D T T Y R

FIG. 5A



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590 610  
gtatacgattatgaccgctgtaatgaccagggccaaaaagaactctt  
V Y D Y D R C N D Q G Q K R T L

630 650 670  
catatagaaaaagccatggaagtcataacgataccgcatatcgataaa  
H I E K A M E V I T I P H I D K

690 710  
gtgcatacacccggaagtaaaagaagttggtaacgctgagatcattggt  
V H T P E V K E V G N A E I I V

730 750 7  
tatgtgcaatcagattatttctcagtggtacaaatggaagattagcggc  
Y V Q S D Y F S V Y K W K I S G

70 790 810  
cgagctgcttttccctcatatcaaacctatttgctggggagtggtctg  
R A A F P S Y Q T Y L L G S V L

830 850  
agcggatcaggacgaatcataaataatgggtattcagtatgaatgcaat  
S G S G R I I N N G I Q Y E C N

870 890 910  
gcaggctcacactttattctgcctgcgcattttggagaatttacaata  
A G S H F I L P A H F G E F T I

930  
gaaggaacatgtgaattcatgatattctcatcct  
E G T C E F M I S H P

**FIG.\_5B**

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10 30  
atgacgcaatcacgatttttctaacgcctgtgtttaagaaaaaatc  
M T Q S P I F L T P V F K E K I

50 70 90  
tggggcggaaccgctttacgagatagatttggatacagtattccttca  
W G G T A L R D R F G Y S I P S

110 130  
gaatcaacgggggaatgctggggccatttccgctcatccaaaaggaccg  
E S T G E C W A I S A H P K G P

150 170 190  
agcactgttgcaaattggcccgtataaaggaaagacattgatcgagctt  
S T V A N G P Y K G K T L I E L

210 230  
tgggaagagcacccgtgaagtattcggcgggcgtagagggggatcggttt  
W E E H R E V F G G V E G D R F

250 270 2  
ccgcttctgacaaagctgctggatgtgaaggaagatacgtcaatttaa  
P L L T K L L D V K E D T S I K

90 310 330  
gttcaccctgatgattactatgccggagaaaacgaagaggggagaactc  
V H P D D Y Y A G E N E E G E L

350 370  
ggcaagacggaatgctggtacattatcgactgtaaggaaaacgcagaa  
G K T E C W Y I I D C K E N A E

390 410 430  
atcatttacgggcatacggcccgtcaaaaaaccgaacttgtcacaatg  
I I Y G H T A R S K T E L V T M

450 470  
atcaacagcgggtgactgggagggcctgctgccaagaatcaaaatttaa  
I N S G D W E G L L R R I K I K

490 510 5  
ccgggtgatttctattatgtgccgagcgggaacgctgcacgcattgtgc  
P G D F Y Y V P S G T L H A L C

30 550 570  
aagggggcccttgtttttagagactcagcaaaattcagatgccacatac  
K G A L V L E T Q Q N S D A T Y

FIG. 6A

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590 610  
cgggtgtacgattatgaccgtcttgatagcaacggaagtccgagagag  
R V Y D Y D R L D S N G S P R E

630 650 670  
cttcatttttgccaaagcgggtcaatgccgccacgggttccccatgtggac  
L H F A K A V N A A T V P H V D

690 710  
gggtatatagatgaatcgacagaatcaagaaaaggaataaccattaaa  
G Y I D E S T E S R K G I T I K

730 750 7  
acatttgtccaagggggaatatatttttcggtttataaatgggacatcaat  
T F V Q G E Y F S V Y K W D I N

70 790 810  
ggcgaagctgaaatgggtcaggatgaatcctttctgatttgcagcgtg  
G E A E M A Q D E S F L I C S V

830 850  
atagaaggaagcggtttgctcaagtatgaggacaaaacatgtccgctc  
I E G S G L L K Y E D K T C P L

870 890 910  
aaaaaagggtgatcacttttattttgccgggtcaaatagcccgattttacg  
K K G D H F I L P A Q M P D F T

930  
ataaaaaggaacttgtacccttatcgtgtctcatatt  
I K G T C T L I V S H I

**FIG.\_6B**